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Search History

Today's Date: 11/30/2000

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USPT	14 near25 cassette	1	<u>L5</u>	
USPT	11 near10 (double-stranded or doublestranded or double stranded)	1072	<u>L4</u>	
USPT	11 near10 masked	3	<u>L3</u>	
USPT	11 and masked	305	<u>L2</u>	
USPT	antisense or anti-sense	9733	<u>L1</u>	

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compns. comprise an antisense masked expression cassette which

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comprise a double-stranded nucleotide sequence and expresses a gene product only in the presence of a target mol. A strand comprises

an

armed expression cassette, i.e., an RNA mol. which codes for a protein of interest linked downstream of a flanking sequence and a translation initiation site operably inserted upstream of the RNA sequence. The flanking sequence encodes a target mol. I.e., the flanking sequence encodes a target get or codes for RNA of interest. The flanking sequence corresponds to the sense strand of the target. A second nucleotide strand

is also provided, capable of hybridizing to the flanking sequence of the first nucleotide sequence, i.e., the antisense strand. The antisense strand masks the translation initiation site when bound. In the presence of a target nucleotide mol., the antisense strand will disassoc. from the armed strand and pair with the target. Dissocn. of the antisense strand unmasks the ribosome binding site allowing the armed cassette to be translated in the presence of the target. A 7-methylguanine cap is used to increase the efficiency of translation. The cassettes are useful for the treatment of disease and for preventing the proliferation of neoplastic cells. Following the protocols, a targeted cassette is constructed wherein the first strand

has

an RNA encoding for toxin A linked with upstream DNA sequences coding the sense portion of the p53 DNA sequence. Inserted within the p53 mol. is a Kozak sequence, and an **antisense** structure is constructed which corresponds to the p53 sense nucleotides.

=> d

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ANSWER 1 OF 1 CAPLUS COPYRIGHT 2000 ACS
    1999:27842 CAPLUS
AN
DN
    130:91265
    RNA vector cassettes for activating and expressing target genes
TI
IN
    Black, Charles Allen, Jr.
PA
SO
    PCT Int. Appl., 43 pp.
    CODEN: PIXXD2
DT
    Patent
    English
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    PATENT NO. KIND DATE APPLICATION NO. DATE
    WO 9858944 A1 19981230 WO 1998-US13093 19980624
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            IE, FI
PRAI US 1997-50772
                     19970625
    WO 1998-US13093 19980624
RE.CNT 2
(1) Coleman; Cell 1984, V37, P429 CAPLUS
(2) Hirashima; Proceedings of the National Academy of Sciences 1986, V83,
P7726
   CAPLUS
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- => s antisense or and sense
- L5 83905 ANTISENSE OR ANTI-SENSE
- => s 15 (10a) masked
- L6 19 L5 (10A) MASKED
- => d 16 ti
- L6 ANSWER 1 OF 19 MEDLINE
- TI In vivo antisense oligodeoxynucleotide mapping reveals masked regulatory elements in an mRNA dormant in mouse occytes.
- => d 2-19 ti
- L6 ANSWER 2 OF 19 MEDLINE
- ${\tt TI}$ Facilitator oligonucleotides increase ribozyme RNA binding to full-length RNA substrates in vitro.
- L6 ANSWER 3 OF 19 MEDLINE
- TI Intranigral administration of D2 dopamine receptor antisense oligodeoxynucleotides establishes a role for nigrostriatal D2 autoreceptors in the motor actions of cocaine.
- L6 ANSWER 4 OF 19 BIOTECHNO COPYRIGHT 2000 Elsevier Science B.V.
- TI In vivo antisense oligodeoxynucleotide mapping reveals masked regulatory elements in an mRNA dormant in mouse oocytes
- L6 ANSWER 5 OF 19 BIOTECHNO COPYRIGHT 2000 Elsevier Science B.V.
- TI Facilitator oligonucleotides increase ribozyme RNA binding to full-length
- RNA substrates in vitro
- L6 ANSWER 6 OF 19 BIOTECHNO COPYRIGHT 2000 Elsevier Science B.V.
- TI Intranigral administration of D.sub.2 dopamine receptor antisense oligodeoxynucleotides establishes a role for nigrostriatal D.sub.2 autoreceptors in the motor actions of cocaine
- L6 ANSWER 7 OF 19 CAPLUS COPYRIGHT 2000 ACS
- TI Masked antisense: a molecular configuration for discriminating similar RNA targets
- L6 ANSWER 8 OF 19 CAPLUS COPYRIGHT 2000 ACS
- TI RNA vector cassettes for activating and expressing target genes
- L6 ANSWER 9 OF 19 CAPLUS COPYRIGHT 2000 ACS
- TI In vivo antisense oligodeoxynucleotide mapping reveals masked regulatory elements in an mRNA dormant in mouse oocytes
- L6 ANSWER 10 OF 19 CAPLUS COPYRIGHT 2000 ACS
- TI Intranigral administration of D2 dopamine receptor antisense oligodeoxynucleotides establishes a role for nigrostriatal D2 autoreceptors in the motor actions of cocaine
- L6 ANSWER 11 OF 19 EMBASE COPYRIGHT 2000 ELSEVIER SCI. B.V.
- TI In vivo antisense oligodeoxynucleotide mapping reveals masked regulatory elements in an mRNA dormant in mouse occytes.
- L6 ANSWER 12 OF 19 EMBASE COPYRIGHT 2000 ELSEVIER SCI. B.V.

- TI Facilitator oligonucleotides increase ribozyme RNA binding to full-length RNA substrates vitro.
- L6 ANSWER 13 OF 19 EMBASE COPYRIGHT 2000 ELSEVIER SCI. B.V.
- TI Intranigral administration of D2 dopamine receptor antisense oligodeoxynucleotides establishes a role for nigrostriatal D2 autoreceptors in the motor actions of cocaine.
- L6 ANSWER 14 OF 19 SCISEARCH COPYRIGHT 2000 ISI (R)
- TI In vivo antisense oligodeoxynucleotide mapping reveals masked regulatory elements in an mRNA dormant in mouse oocytes
- L6 ANSWER 15 OF 19 SCISEARCH COPYRIGHT 2000 ISI (R)
- TI FACILITATOR OLIGONUCLEOTIDES INCREASE RIBOZYME RNA-BINDING TO FULL-LENGTH RNA SUBSTRATES IN-VITRO
- L6 ANSWER 16 OF 19 SCISEARCH COPYRIGHT 2000 ISI (R)
- TI INTRANIGRAL ADMINISTRATION OF D-2, DOPAMINE-RECEPTOR ANTISENSE OLIGODEOXYNUCLEOTIDES ESTABLISHES A ROLE FOR NIGROSTRIATAL D-2 AUTORECEPTORS IN THE MOTOR ACTIONS OF COCAINE
- L6 ANSWER 17 OF 19 BIOSIS COPYRIGHT 2000 BIOSIS
- TI In vivo antisense oligodeoxynucleotide mapping reveals masked regulatory elements in an mRNA dormant in mouse oocytes.
- L6 ANSWER 18 OF 19 BIOSIS COPYRIGHT 2000 BIOSIS
- TI Facilitator oligonucleotides increase ribozyme RNA binding to full-length RNA substrates in vitro.
- L6 ANSWER 19 OF 19 BIOSIS COPYRIGHT 2000 BIOSIS
- TI Intranigral administration of D-2 dopamine receptor antisense oligodeoxynucleotides establishes a role for nigrostriatal D-2 autoreceptors in the motor actions of cocaine.
- => d ab 5 7 8 12 13
- ANSWER 5 OF 19 BIOTECHNO COPYRIGHT 2000 Elsevier Science B.V.

 Primer extension arrest (PEA) studies have demonstrated that antisense oligonucleotides (.beta.112C, .beta.114C), which lie upstream of a ribozyme targeted to .beta.-amyloid peptide precursor (.beta.APP) mRNA, but not sense oligonucleotides (.beta.112S, .beta.116S) or a scrambled oligonucleotide, .beta.116M, affect ribozyme-mediated cleavage in vitro. Substrate dissociation experiments revealed that the ribozyme binding site in this mRNA was masked; PEA kinetics showed the association of the ribozyme and substrate was enhanced by antisense oligonucleotide binding. These studies suggest that masked ribozyme cleavage sites that may occur in disease-causing mRNAs can be targeted for degradation using 'facilitator' oligonucleotides.
- L6 ANSWER 7 OF 19 CAPLUS COPYRIGHT 2000 ACS
- AB Antisense technol. has great potential for the control of RNA expression, but there remain few successful applications of the technol. Expressed antisense RNA can effectively down-regulate expression of a gene over
- periods, but cannot differentiate partly identical sequences, such as the mRNA of fusion genes or those with point mutants. We have designed a structured form of expressed antisense, which can discriminate between highly similar mRNA mols. These 'masked' antisense RNAs have most of the antisense sequence sequestered within duplex elements, leaving a short single-stranded region to initiate binding to target RNA. After contacting the correct target, the structured RNA can unravel, releasing the masked antisense region to form a stable duplex with the mRNA. We

demonstrate that suitable masked antisense RNA can discriminate between the two forms of BCR-ABL mast that result from the Philadelphia chamosomal translocations, as well as discriminating the normal BCR and ABl mRNA.

L6 ANSWER 8 OF 19 CAPLUS COPYRIGHT 2000 ACS

AB Compns. and methods for activating genes of interest are provided. The compns. comprise an antisense masked expression cassette which comprise a double-stranded nucleotide sequence and expresses a gene product only in the presence of a target mol. A first strand comprises an armed expression cassette, i.e., an RNA mol. which codes for a protein of interest linked downstream of a flanking sequence and a translation initiation site operably inserted upstream of the RNA sequence. The flanking sequence encodes a target mol. I.e., the

flanking
sequence encodes a target get or codes for RNA of interest. The flanking
sequence corresponds to the sense strand of the target. A second
nucleotide strand is also provided, capable of hybridizing to the

flanking

sequence of the first nucleotide sequence, i.e., the antisense strand. The antisense strand masks the translation initiation site when bound.

Ιn

the presence of a target nucleotide mol., the antisense strand will disassoc. from the armed strand and pair with the target. Dissocn. of

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antisense strand unmasks the ribosome binding site allowing the armed cassette to be translated in the presence of the target. A 7-methylguanine cap is used to increase the efficiency of translation. The cassettes are useful for the treatment of disease and for preventing the proliferation of neoplastic cells. Following the protocols, a targeted cassette is constructed wherein the first strand has an RNA encoding for toxin A linked with upstream DNA sequences coding the sense portion of the p53 DNA sequence. Inserted within the p53 mol. is a Kozak sequence, and an antisense structure is constructed which corresponds to the p53 sense nucleotides.

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AB Primer extension arrest (PEA) studies have demonstrated that antisense oligonucleotides (.beta.112C, .beta.114C), which lie upstream of a ribozyme targeted to .beta.-amyloid peptide precursor (.beta.APP) mRNA, but not sense oligonucleotides (.beta.112S, .beta.116S) or a scrambled oligonucleotide, .beta.116M, affect ribozyme-mediated cleavage in vitro. Substrate dissociation experiments revealed that the ribozyme binding

site

in this mRNA was masked; PEA kinetics showed the association of the ribozyme and substrate was enhanced by **antisense** oligonucleotide binding. These studies suggest that **masked** ribozyme cleavage sites that may occur in disease-causing mRNAs can be targeted for degradation using 'facilitator' oligonucleotides.

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Dopamine D2 autoreceptors found on nigrostriatal dopaminergic neurons are thought to inhibit dopamine release, tyrosine hydroxylase activation, and spontaneous firing rate. It is likely that these receptors play an important role in moderating the behavioral response to cocaine, but the lack of potent selective autoreceptor ligands has made it difficult to assess this contribution. We have developed an antisense phosphorothicate oligodeoxynucleotide (ODN) against D2 receptor mRNA, which was used to reduce levels of D2 receptors in vitro and in vivo. Unilateral administration of antisense ODN, via intracerebral cannula, into the substantia nigra of rats for several days caused dramatic contralateral rotational behavior in response to a subcutaneous injection of cocaine. This effect was maximal by 10 min after injection of cocaine and lasted for >30 min; without cocaine, no spontaneous rotational behavior was noted. In striatal slices, the potency of sulpiride, a D2 antagonist, in

enhancing electrically stimulated dopamine release was significantly reduced on the isense-treated side; this is distent with a decrease in the striatal 2 autoreceptor population. As measured by quantitative autoradiography, administration of antisense ODN caused a loss of approximately 40% of nigral D2 receptor [125I]iodosulpride binding, compared with the untreated side. In vitro, treatment of WERI-27 retinoblastoma cells with D2 antisense ODN at a concentration of 1 .mu.M reduced D2 receptor levels by 57% after 3 days. The robustness of cocaine-induced rotation and the impaired ability of sulpiride to enhance dopamine release from slices suggest that nigrostriatal D2 autoreceptors play a direct role in reducing the motor response to cocaine administration. Furthermore, the absence of spontaneous rotation in antisense ODN-treated animals suggests that autoreceptor effects are masked by compensatory mechanisms during normal behavior.

=> d 5 7 8 12 cit

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- ANSWER 5 OF 19 BIOTECHNO COPYRIGHT 2000 Elsevier Science B.V. L6 1996:26092241 ANBIOTECHNO Facilitator oligonucleotides increase ribozyme RNA binding to TIfull-length RNA substrates in vitro ΑU Denman R.B. Department of Molecular Biology, New York State Institute, Basic CS Res. Developmental Disabilities, 1050 Forest Hill Road, Staten Island, NY 10314, United States. FEBS Letters, (1996), 382/1-2 (116-120) SO CODEN: FEBLAL ISSN: 0014-5793 DT Journal; Article ' Netherlands CYLΑ English English \mathtt{SL} ANSWER 7 OF 19 CAPLUS COPYRIGHT 2000 ACS L62000:639398 CAPLUS ΑN TIMasked antisense: a molecular configuration for discriminating similar RNA targets Stocks, Martin R.; Rabbitts, Terence H. ΑU MRC Laboratory Molecular Biology, Cambridge, CB2 2QH, UK CŚ EMBO Rep. (2000), 1(1), 59-64 SO CODEN: ERMEAX; ISSN: 1469-221X Oxford University Press PΒ DTJournal English LΑ RE.CNT 20
- (1) Agrawal, S; Trends Biotechnol 1996, V14, P376 CAPLUS
- (2) Ayub, R; Nature Biotechnol 1996, V14, P862 CAPLUS
- (3) Bartram, C; Nature 1983, V306, P277 CAPLUS
- (4) de Klein, A; Nature 1982, V300, P765 CAPLUS
- (6) Groffen, J; Cell 1984, V36, P93 CAPLUS
- ALL CITATIONS AVAILABLE IN THE RE FORMAT

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L6
    ANSWER 8 OF 19 CAPLUS COPYRIGHT 2000 ACS
    1999:27842 CAR
ΑN
DN
    130:91265
    RNA vector cassettes for activating and expressing target genes
TI
IN
    Black, Charles Allen, Jr.
PA
SO
     PCT Int. Appl., 43 pp.
     CODEN: PIXXD2
DT
    Patent
LA
    English
FAN.CNT 1
                                        APPLICATION NO. DATE
     PATENT NO.
                    KIND DATE
     WO 1998-US13093 19980624
    WO 9858944
PI
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RE.CNT 2
(1) Coleman; Cell 1984, V37, P429 CAPLUS
(2) Hirashima; Proceedings of the National Academy of Sciences 1986, V83,
P7726
   CAPLUS
    ANSWER 12 OF 19 EMBASE COPYRIGHT 2000 ELSEVIER SCI. B.V.
L6
ΑN
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TΙ
     Facilitator oligonucleotides increase ribozyme RNA binding to full-length
    RNA substrates in vitro.
ΑU
    Denman R.B.
    Department of Molecular Biology, New York State Institute, Basic
CS
    Res. Developmental Disabilities, 1050 Forest Hill Road, Staten Island, NY
     10314, United States
    FEBS Letters, (1996) 382/1-2 (116-120).
SO
    ISSN: 0014-5793 CODEN: FEBLAL
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            Pharmacology
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